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Substitute for form 1449/PTO				Complete if Known	
INFORMATION DISCLOSURE STATEMENT BY APPLICANT				Application Number	10/048033
Date Submitted: August 30, 2007 <i>(use as many sheets as necessary)</i>				Filing Date	7/21/2000
				First Named Inventor	H. Michael Shepard
				Art Unit	1623
				Examiner Name	Crane, Lawrence E.
				Attorney Docket Number	060925-0601
Sheet	1	of	5		

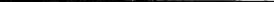
U.S. PATENT DOCUMENTS

UNPUBLISHED U.S. PATENT APPLICATION DOCUMENTS

SEARCHED, SERIALIZED, AND INDEXED PATENT APPLICATION DOCUMENTS					
Examiner Initials*	Cite No. ¹	U.S. Patent Application Document	Filing Date of Cited Document MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Serial Number-Kind Code ² (if known)			

FOREIGN PATENT DOCUMENTS

Examiner Initials*	Cite No. ¹	Foreign Patent Document	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Documents	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T- ⁶
		Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)				
	F15	WO 96/23506	08-08-1996	Fraunhofer Society for the Promotion of Applied Research E.V. (English Abstract)		

Examiner Signature	L. E. Crane 	Date Considered	11/26/2007
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	D16	BALZARINI et al. "Highly Selective Cytostatic Activity of (E)-5-(2-Bromovinyl)-2'-deoxyuridine Derivatives for Murine Mammary Carcinoma (FM3A) Cells Transformed with the Herpes Simplex Virus Type 1 Thymidine Kinase Gene" Molecular Pharmacology (1985) 28:581-587	
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	D19	BATHE et al. "Increased thymidylate synthase gene expression in liver metastases from colorectal carcinoma: implications for chemotherapeutic options and survival" Cancer J. Sci. Am. 5(1):34-40 (1999)	
	D20	BIBLE et al. "Cytotoxic synergy between flavopiridol (NSC 649890, L86-8275) and various antineoplastic agents: the importance of sequence of administration" Cancer Res. 57(16):3375-80 (August 15, 1997)	
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	D22	CASS et al. "Recent advances in the molecular biology of nucleoside transporters of mammalian cells" Biochem. Cell Biol. 76(5):761-770 (1998)	
	D23	CHOU and TALALAY, "Quantitative analysis of dose-effect relationships: the combined effects of multiple drugs or enzyme inhibitors" Adv. Enzyme Regul. 22:27-55 (1984)	
	D24	CORDON-CARDO and PRIVES "At the crossroads of inflammation and tumorigenesis" J. Exp. Med. 190(10):1367-1370 (November 15, 1999)	
	D25	COSTI et al. "Phthalein derivatives as a new tool for selectivity in thymidylate synthase inhibition" J. Med. Chem. 42(12):2112-2124 (1999)	
	D26	COSTI, et al. "Thymidylate Synthase Inhibition: A Structure-Based Rationale for Drug Design" Medical Research Review 18(1):21-42 (1998)	
	D27	CURT "Cancer drug development: new targets for cancer treatment" Oncologist 1(3):II-III (1996)	

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Sheet	3	of	5	Attorney Docket Number	060925-0601

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published.			
	D28	DeCLERCQ et al. "Antiviral Activity of Novel Deoxyuridine Derivatives" Current Chemotherapy: Proceedings of the International Congress of Chemotherapy 1:352-354 (September 18, 1978)			
	D29	DeCLERCQ "Antiviral Activity Spectrum and Target of Action of Different Classes of Nucleoside Analogues" Nucleosides & Nucleosides 13(6&7):1271-1295 (1994)			
	D30	DeCLERCQ "In search of a selective antiviral chemotherapy" Clin. Micro. Review 10(4):674-693 (October 1997)			
	D31	DRAKE et al. "Resistance to Tomudex (ZD1694): Multifactorial in Human Breast and Colon Carcinoma Cell Lines" Biochem. Pharmacol. 51(10):1349-1355 (1996)			
	D32	FARROW et al. "Synthesis and biological properties of novel phosphotriesters: A new approach to the introduction of biologically active nucleotides into cells" J. Med. Chem. 33(5):1400-1406 (1990)			
	D33	GREM "Biochemical modulation of fluorouracil by dipyridamole: preclinical and clinical experience" Semin Oncol. 19(2 Suppl 3):56-65 (April 1992)			
	D34	GRIENGL et al. "Phosphonoformate and phosphonoacetate derivatives of 5-substituted 2'deoxyuridines: Synthesis and antiviral activity" J. Med. Chem. 31(9):1831-1839 (1988)			
	D35	HENGSTSCHLAGER, M. and E. WAWRA "Cytofluorometric determination of thymidine kinase activity in a mixture of normal and neoplastic cells" Br. J. Cancer (1993) 67:1022-1025			
	D36	JOHNSTON et al. "Thymidylate synthase gene and protein expression correlate and are associated with response to 5-fluorouracil in human colorectal and gastric tumors" Cancer Res. 55:1407-1412 (1995)			
	D37	KOMAKI et al. "Difference in thymidylate synthetase activity in involved nodes compared with primary tumor in breast cancer patients" Breast Cancer Res. Treat. 35(2):157-162 (1995)			
	D38	LEICHMAN "Thymidylate synthase as a predictor of response" Oncol. 12(8 Suppl 6):43-47 (August 1998)			
	D39	LIBRA, et al. "Thymidylate synthase mRNA levels are increased in liver metastases of colorectal cancer patients resistant to fluoropyrimidine-based chemotherapy" BMC Cancer 4:11 (2004)			
	D40	LIVINGSTON et al. "Studies with tetrahydrofolate and thymidylate synthetase from amethopterin-resistant mouse leukemia cells" Biochem. 7(8):2814-2818 (1968)			

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				Art Unit	1623
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	D41	LOOK et al. "Increased thymidine kinase and thymidylate synthase activities in human epithelial ovarian carcinoma" <i>Anticancer Res.</i> 17:2353-2356 (1997)		
	D42	MIAO et al. "A Stepwise One Pot Synthesis of Alkyl Thiophosphoramidate Derivatives of Nucleosides" <i>Synthetic Communications</i> 32(8):1159-1167 (2002)		
	D43	NICULESCU-DUVAZ et al. "Gene-directed enzyme prodrug therapy: A review of enzyme/prodrug combinations" <i>Expert Opin. Invest. Drugs</i> 6(6):685-703 (1997)		
	D44	PATTERSON et al. "Thymidine phosphorylase moderates thymidine-dependent rescue after exposure to the thymidylate synthase inhibitor ZD1694 (tomudex) in vitro" <i>Cancer Res.</i> 58:2737-2740 (1998)		
	D45	PEDERSEN-LANE et al. "High-level expression of human thymidylate synthase" <i>Protein Expression and Purification</i> 10:256-262 (1997)		
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	D47	SMITH et al. "Response to doxorubicin of cultured normal and cancerous human mammary epithelial cells" <i>J. Nat'l Cancer Inst.</i> 74(2):341-347 (1985)		
	D48	TOUROUTOGLOU et al. "Thymidylate synthase inhibitors" <i>Clin. Cancer Res.</i> 2(2):227-243 (February 1996)		
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	D51	van TRIEST et al. "Thymidylate synthase level as the main predictive parameter for sensitivity to 5-fluorouracil, but not for folate-based thymidylate synthase inhibitors, in 13 nonselected colon cancer lines" <i>Clin. Cancer Res.</i> 5(3):643-654 (1999)		
	D52	VLAYKAVA et al. "Increased Thymidylate Synthase Gene Expression in Metastatic Melanoma" <i>Oncology</i> 54:146-152 (1997)		
	D53	WHALEN et al. "Human glutathione S-transferases" <i>Seminars in Liver Disease</i> 18(4):345-358 (1998)		

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<i>JRC</i>	D54	WILDNER et al. "Enzyme prodrug gene therapy: Synergistic use of the herpes simplex virus-cellular thymidine kinase/ganciclovir system and thymidylate synthase inhibitors for the treatment of colon cancer" n Cancer Res. 59(20):5233-5238 (1999)	
<i>JRC</i>	D55	WILLSON et al. "Phase I Clinical Trial of a Combination of Dipyridamole and Acivicin Based Upon Inhibition of Nucleoside Salvage" Cancer Res. 48:5585-5590 (October 1, 1998)	
<i>JRC</i>	D56	WOLFE et al. "Antibody-directed enzyme prodrug therapy with the T268G mutant of human carboxypeptidase A1: in vitro and in vivo studies with prodrugs of methotrexate and the thymidylate synthase" Bioconjugate Chemistry 10(1):38-48 (1999)	

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